

Amendments to the Claims :

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (Currently amended) Apparatus for electroporation comprising a wave generator, a biochip containing an array of microelectrodes and a control system that permits to transfer ~~the~~ of a signal to a pre-selected single microelectrode of the biochip.
2. (Currently amended) Apparatus according to claim 1 ~~characterised in that~~ wherein the control system consists of a personal computer equipped with a software program capable of designing various waveform signals and a switching system controlling the wave generator output.
3. (Currently amended) Apparatus according to claims 1 ~~and 2 characterised in that~~ wherein the biochip comprises an array of microelectrodes of a size comparable to a ~~the~~ cell to be electroporated and each of said microelectrodes being driven separately from the others allowing very precise and punctual control of the electroporation process.
4. (Currently amended) A biochip ~~Biochip~~ comprising an array of individually driven microelectrodes ~~(20)~~ comprised on a suitable insulating layer mounted on a solid substrate; means to electrically connect said microelectrodes to a switching system; a cell culture chamber where ~~the~~ cells can be grown and adhere in contact with said array of microelectrodes on a surface formed by said insulating layer containing said array of microelectrodes on said solid substrate.
5. (Currently amended) The biochip ~~Biochip~~ according to claim 4 comprising a semiconductor substrate as the solid substrate covered with an insulating layer ~~(27)~~ comprising said array of individually driven microelectrodes ~~(20)~~ of a size comparable to the cell to be electroporated, and mounting a cell culture chamber ~~(24)~~ with an opening ~~(26)~~ mounted, in turn, on a support ~~(21)~~ made of dielectric

material, said microelectrodes (20) being electrically connected via conductive traces (28) to conductive pads (29) electrically connected, in turn, to a couple of external parallel connectors (22) through wire bonding (23) covered by an ~~the~~ outer portion of the cell culture chamber (24) encircling the opening (26), being said cell culture chamber (24) with the opening (26) mounted over the top of ~~the~~ said semiconductor substrate covered with the ~~an~~ insulating layer (27), both attached on the dielectric support (21).

6. (Currently amended) A biochip ~~Biochip~~ according to claim 5 comprising two further electrodes (25) integrated in the semiconductor substrate covered with the ~~an~~ insulating layer (27), and acting as a ground reference.
7. (Currently amended) A biochip ~~Biochip~~ according to claim 5 wherein the semiconductor substrate covered with the ~~an~~ insulating layer (27) is a silicon substrate covered with an ~~a~~ insulating layer preferentially of SiO₂.
8. (Currently amended) The biochip ~~Biochip~~ according to claim 5 wherein the ~~these~~ solid substrates are is transparent.
9. (Currently amended) The biochip ~~Biochip~~ according to claim 5 wherein the dielectric support is vetronite, glass or ceramic.
10. (Currently amended) The biochip ~~Biochip~~ according to claim 5 wherein the microelectrodes of the array (20) have a size with a surface of at least ten per cent of the total cell membrane and preferably a diameter ranging from 1 μ m to 50 μ m.
11. (Currently amended) The biochip ~~Biochip~~ according to claims 4 –10 wherein the microelectrodes are of conductive or capacitive type.
12. (Currently amended) The microelectrodes ~~Microelectrodes~~ according to claim 11 consisting of conductive microelectrodes obtained over a silicon substrate (31) covered with an ~~a~~ insulating layer preferentially of SiO₂ (32), said microelectrodes having connecting traces wherein said microelectrodes and their connecting traces

(38) being made by a "sandwich" of two titanium nitride, (TiN), layers (33) and an aluminium layer (34), covered with a gold layer (37) on their active surface.

13. (Currently amended) The microelectrodes ~~Microelectrodes~~ according to claim 11 wherein said microelectrodes are realized ~~realised~~ using Metal Oxide Semiconductor, (MOS), technology.
14. (Currently amended) The microelectrodes ~~Microelectrodes~~ according to claim 13 consisting of a silicon p-type substrate (40) in which two n-doped regions, a drain (41) and a source (42), are implanted with conventional microelectronic techniques, ~~the~~ a gate (43) of these electrodes being realized ~~realised~~ in n+ doped polysilicon and is common to all devices in a row, (word line), the drain (41) of all devices in a column being connected together by using a metal contact plug and a metal line (44), the source (42) of the transistor being connected via a metal, (usually tungsten), plug (46) to a gold layer (47) which acts as the active electrode.
15. (Currently amended) The microelectrodes ~~Microelectrodes~~ according to claim 11 consisting of a capacitive microelectrode obtained with an insulating substrate (60), a metal (61) and a thin insulating layer (64) said microelectrodes being separated by insulating material (62) and covered in non exposed areas by a passivation layer (63).
16. (Currently amended) A method ~~Method~~ of electroporation ~~characterised in that~~ wherein an apparatus according to claims 1 —3 is used.
17. (Currently amended) The method ~~Method~~ according to claim 16 ~~characterised in that~~ wherein said apparatus performs one or more electroporations to at least a single adhering cell.
18. (Currently amended) The method ~~Method~~ according to claim 17 ~~characterised in that~~ wherein said apparatus comprises a biochip according to claims 4 —11.
19. (Currently amended) The method ~~Method~~ according to claims 16 —18 ~~characterised~~

~~in that wherein~~ said biochip comprises microelectrodes according to claims ~~11 12—~~ 15.

20. (Currently amended) ~~The method Method~~ according to claims 16 –~~19~~ characterised ~~in that wherein~~ the wave generator sends to the electrodes trains of pulses of various amplitude and duration.

21. (Currently amended) ~~The method Method~~ according to claims 16— 20 characterised ~~in that wherein~~ the wave generator sends to the microelectrodes five trains of 25 pulses, {1 ms duration}, repeated at a time interval of 500 ms.

22. (Currently amended) ~~The method Method~~ according to claims 16—20 characterised ~~in that wherein~~ a trains of triangular voltages consisting of 10 pulses are applied to the electrodes the interval between one train and another being of 5 s.

23. (Currently amended) ~~The method Method~~ according to ~~any of preceding~~ claims 16 characterised ~~in that it comprises~~ substantially the following steps:

- ~~cultivating~~ cultivate cells since the adhering stage is reached;
- ~~adding~~ add in the culture medium at least one compound to be electroporated in at least one single cell of the said cells;
- ~~selecting~~ selected at least one single cell and at least one microelectrode on which said selected single cell is adherent;
- ~~generating~~ generate at least one electric signal ~~suitable~~ capable to electroporate said at least one single cell with said at least one compound to be electroporated and ~~driving~~ drive said electric signal to the said one microelectrode on which said selected single cell is adherent.

24. (Canceled) ~~Electroporated cells characterised in that they are obtained with method according claims 16—23.~~

25. (Canceled) ~~Electroporated cells according to claim 24 wherein the electroporated agents are drugs, genetic constructs and proteins.~~

26. (New) The method Method according to claim 23 wherein the said compounds to be electroporated are drugs, genetic constructs and proteins.

